- A method for identifying a modulator of the enzymatic production of β-amyloid peptide (Aβ) from β-amyloid precursor protein (APP) or a fragment thereof, comprising:
 providing a APP protease and BACE2 to a sample containing APP or an APP fragment;
 contacting the APP or the APP fragment and BACE2 with a candidate compound; and
 monitoring the effect of the candidate compound on the production of Aβ.
 - 2. The method of claim 1, wherein said modulator is an inhibitor of Aβ production.
- 3. The method of claim 1, wherein the APP protease is a protease with α -secretase activity.
- 4. The method of claim 1, wherein the APP protease is a protease with a γ -secretase activity.
- 5. The method of claim 1, wherein the APP protease is a protease with a β -secretase activity other than BACE2.
- 6. The method of claim 5, wherein said β -secretase activity is due to the presence of an enzyme having a pH optimum at about pH 6.5-7.0, and an estimated molecular weight of about 32-39 kDa as calculated from radiation inactivation analysis of HEK293 cell membrane extracts, or about 20-26 kDa as calculated from radiation inactivation analysis of human brain samples, with a candidate compound.
- 7. The method of claim 5, wherein said β-secretase activity is due to the presence of a β-secretase enzyme having a pH optimum at about pH 4.5-5.0 and an estimated molecular weight of about 50-60 kDa as calculated from radiation inactivation analysis of HEK293 cell membrane extracts or human brain samples (BACEI).
- 8. The method of claim 1, wherein the effect of the candidate compound on the production of A β is monitored by measuring the amount of A β formed.

- 9. The method of claim 8, wherein the amount of $A\beta$ formed is reduced by at least about 50%.
- 10. The method of claim 8, wherein the amount of $A\beta$ formed is reduced by at least about 75%.
- 11. The method of claim 8, wherein the amount of $A\beta$ formed is reduced by at least about 90%.
- 12. The method of claim 1, further comprising the step of comparing the effect of the test compound on A β production with the effect of BACE2 in the absence of the test compound.
- 13. The method of claim 12, wherein the test compound causes at least about 15% reduction in the amount of A β over the effect of BACE2 in the absence of the test compound.
 - 14. The method of claim 13, wherein said reduction is at least about 25%.
 - 15. The method of claim 13, wherein said reduction is at least about 50%.
 - 16. The method of claim 13, wherein said reduction is at least about 75%
- 17. The method of claim 1, further comprising the step of comparing the effect of the test compound on A β production with A β production in the absence of BACE2.
 - 18. The method of claim 1, which is performed in a cell-free format.
- 19. A method for reducing the amount of β -amyloid deposits in the central nervous system (CNS) of a mammal comprising administering to said mammal an effective amount o of BACE2 or an agonist thereof.
 - 20. The method of claim 19 wherein said mammal is human.